

10/586124

IAP11 Rec'd PCT/PTO 14 JUL 2006

5189.ST25.txt  
SEQUENCE LISTING

<110> Bayer Pharmaceuticals Corporation  
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Lumb, Kevin  
Buckholz, Thomas  
Salhanick, Arthur

<120> PITUITARY ADENYLYL CYCLASE ACTIVATING PEPTIDE (PACAP) RECEPTOR  
(VPAC2) AGONISTS AND THEIR PHARMACOLOGICAL METHODS OF USE

<130> 5189

<150> US 60/539,550  
<151> 2004-01-27

<150> US 60/566,499  
<151> 2004-04-29

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<170> PatentIn version 3.3

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<400> 61

His Ser Asp Ala Val Phe Thr Asp Asn Tyr Thr Arg Leu Arg Lys Gln  
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Met Ala Ala Lys Lys Tyr Leu Gln Ser Ile Lys Gln Lys Leu  
20 25 30

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<400> 62

His Ser Asp Ala Val Phe Thr Asp Asn Tyr Thr Arg Leu Arg Lys Gln  
1 5 10 15

Met Ala Ala Lys Lys Tyr Leu Gln Ser Ile Lys Gln Lys Met  
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<400> 63

His Ser Asp Ala Val Phe Thr Asp Asn Tyr Thr Arg Leu Arg Lys Gln  
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Met Ala Ala Lys Lys Tyr Leu Gln Ser Ile Lys Gln Lys Pro  
20 25 30

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<400> 64

His Ser Asp Ala Val Phe Thr Asp Asn Tyr Thr Arg Leu Arg Lys Gln  
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Met Ala Ala Lys Lys Tyr Leu Gln Ser Ile Lys Gln Lys Gln  
20 25 30

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<400> 65

His Ser Asp Ala Val Phe Thr Asp Asn Tyr Thr Arg Leu Arg Lys Gln  
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Met Ala Ala Lys Lys Tyr Leu Gln Ser Ile Lys Gln Lys Ser  
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His Ser Asp Ala Val Phe Thr Asp Asn Tyr Thr Arg Leu Arg Lys Gln  
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Met Ala Ala Lys Lys Tyr Leu Gln Ser Ile Lys Gln Lys Thr  
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His Ser Asp Ala Val Phe Thr Asp Asn Tyr Thr Arg Leu Arg Lys Gln  
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Met Ala Ala Lys Lys Tyr Leu Gln Ser Ile Lys Gln Lys Val  
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<210> 68  
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<400> 68

His Ser Asp Ala Val Phe Thr Asp Asn Tyr Thr Arg Leu Arg Lys Gln  
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Met Ala Ala Lys Lys Tyr Leu Gln Ser Ile Lys Gln Lys Trp  
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Met Ala Ala Lys Lys Tyr Leu Gln Ser Ile Lys Gln Lys Tyr  
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His Ser Asp Ala Val Phe Thr Asp Asn Tyr Thr Arg Leu Arg Lys Gln  
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Met Ala Gly Lys Lys Tyr Leu Gln Ser Ile Lys Gln Arg Ile  
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<400> 71

His Ser Asp Ala Val Phe Thr Asp Asn Tyr Thr Arg Leu Arg Lys Gln  
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Met Ala Lys Lys Lys Tyr Leu Gln Ser Ile Lys Gln Arg Ile  
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His Ser Asp Ala Val Phe Thr Asp Asn Tyr Thr Arg Leu Arg Lys Gln  
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Met Ala Ser Lys Lys Tyr Leu Gln Ser Ile Lys Gln Arg Ile  
20 25 30

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His Ser Asp Ala Val Phe Thr Asp Asn Tyr Thr Arg Leu Arg Lys Gln  
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Met Ala Ala Lys Lys Tyr Leu Gln Ser Ile Pro Gln Arg Ile  
20 25 30

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His Ser Asp Ala Val Phe Thr Asp Asn Tyr Thr Arg Leu Arg Lys Gln  
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Met Ala Ser Lys Lys Tyr Leu Gln Ser Ile Arg Gln Arg Ile  
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<400> 75

His Ser Asp Ala Val Phe Thr Asp Gln Tyr Thr Arg Leu Arg Lys Gln  
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Val Ala Ala Lys Lys Tyr Leu Gln Ser Ile Lys Asn Lys Arg Tyr  
20 25 30

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His Ser Asp Ala Val Phe Thr Asp Gln Tyr Thr Arg Leu Arg Lys Gln  
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Met Ala Ala Lys Lys Tyr Leu Gln Ser Ile Lys Asn Lys Arg Tyr  
20 25 30

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<400> 77

His Ser Asp Ala Val Phe Thr Asp Gln Tyr Thr Arg Leu Arg Lys Gln  
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Val Ala Ala Lys Lys Tyr Leu Gln Ser Ile Lys Asn Lys  
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5189.ST25.txt

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<400> 78

His Thr Glu Ala Val Phe Thr Asp Gln Tyr Thr Arg Leu Arg Lys Gln  
1 5 10 15

Val Ala Ala Lys Lys Tyr Leu Gln Ser Ile Lys Asn Lys Arg Tyr  
20 25 30

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<400> 79

His Ser Asp Ala Val Phe Thr Asp Gln Tyr Thr Arg Leu Arg Lys Gln  
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Leu Ala Val Lys Lys Tyr Leu Gln Asp Ile Lys Asn Gly Gly Thr  
20 25 30

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His Ser Asp Ala Val Phe Thr Asp Gln Tyr Thr Arg Leu Arg Lys Gln  
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Met Ala Ala Lys Lys Tyr Leu Gln Ser Ile Lys Asn Lys Arg  
20 25 30

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<400> 81

His Ser Asp Ala Val Phe Thr Asp Gln Tyr Thr Arg Leu Arg Lys Gln  
1 5 10 15

Leu Ala Ala Lys Lys Tyr Leu Gln Thr Ile Lys Asn Lys Arg Tyr  
20 25 30

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<400> 82

His Ser Asp Ala Val Phe Thr Asp Gln Tyr Thr Arg Leu Arg Lys Gln  
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Met Ala Ala Lys Lys Tyr Leu Gln Thr Ile Lys Asn Lys Arg Tyr  
20 25 30

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<400> 83

His Ser Asp Ala Val Phe Thr Asp Gln Tyr Thr Arg Leu Arg Lys Gln  
1 5 10 15

Met Ala Ala His Lys Tyr Leu Gln Ser Ile Lys Asn Lys Arg Tyr  
20 25 30

5189.ST25.txt

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<400> 84

His Ser Asp Ala Val Phe Thr Asp Gln Tyr Thr Arg Leu Arg Lys Gln  
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Met Ala Ala Lys His Tyr Leu Gln Ser Ile Lys Asn Lys Arg Tyr  
20 25 30

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<400> 85

His Ser Asp Ala Val Phe Thr Asp Gln Tyr Thr Arg Leu Arg Lys Gln  
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Met Ala Gly Lys Lys Tyr Leu Gln Ser Ile Lys Asn Lys Arg  
20 25 30

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<400> 86

His Ser Asp Ala Val Phe Thr Asp Gln Tyr Thr Arg Leu Arg Lys Gln  
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Met Ala Lys Lys Lys Tyr Leu Gln Ser Ile Lys Asn Lys Arg  
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<400> 87

His Ser Asp Ala Val Phe Thr Asp Gln Tyr Thr Arg Leu Arg Lys Gln  
1 5 10 15

Met Ala Arg Lys Lys Tyr Leu Gln Ser Ile Lys Asn Lys Arg  
20 25 30

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His Ser Asp Ala Val Phe Thr Asp Gln Tyr Thr Arg Leu Arg Lys Gln  
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Met Ala Ser Lys Lys Tyr Leu Gln Ser Ile Lys Asn Lys Arg  
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<400> 89

His Ser Asp Ala Val Phe Thr Asp Gln Tyr Thr Arg Leu Arg Lys Gln  
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Met Ala Ala Lys Lys Tyr Leu Gln Ser Ile Pro Asn Lys Arg  
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His Ser Asp Ala Val Phe Thr Asp Gln Tyr Thr Arg Leu Arg Lys Gln  
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Met Ala Ala Lys Lys Tyr Leu Gln Ser Ile Gln Asn Lys Arg  
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His Ser Asp Ala Val Phe Thr Asp Gln Tyr Thr Arg Leu Arg Lys Gln  
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Met Ala Ala Lys Lys Tyr Leu Gln Ser Ile Arg Asn Lys Arg  
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<400> 92

His Ser Asp Ala Val Phe Thr Asp Gln Tyr Thr Arg Leu Arg Lys Gln  
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Met Ala Ala Lys Lys Tyr Leu Gln Ser Ile Lys Asn Arg Arg  
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His Ser Asp Ala Val Phe Thr Asp Gln Tyr Thr Arg Leu Arg Lys Gln  
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Met Ala Ala Lys Lys Tyr Leu Gln Ser Ile Lys Asn Lys Ala  
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His Ser Asp Ala Val Phe Thr Asp Gln Tyr Thr Arg Leu Arg Lys Gln  
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Met Ala Ala Lys Lys Tyr Leu Gln Ser Ile Lys Asn Lys Phe  
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<400> 95

His Ser Asp Ala Val Phe Thr Asp Gln Tyr Thr Arg Leu Arg Lys Gln  
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Met Ala Ala Lys Lys Tyr Leu Gln Ser Ile Lys Asn Lys His  
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Met Ala Ala Lys Lys Tyr Leu Gln Ser Ile Lys Asn Lys Ile  
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Met Ala Ala Lys Lys Tyr Leu Gln Ser Ile Lys Asn Lys Lys  
20 25 30

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<400> 98

His Ser Asp Ala Val Phe Thr Asp Gln Tyr Thr Arg Leu Arg Lys Gln  
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Met Ala Ala Lys Lys Tyr Leu Gln Ser Ile Lys Asn Lys Leu  
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His Ser Asp Ala Val Phe Thr Asp Gln Tyr Thr Arg Leu Arg Lys Gln  
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Met Ala Ala Lys Lys Tyr Leu Gln Ser Ile Lys Asn Lys Met  
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<400> 100

His Ser Asp Ala Val Phe Thr Asp Gln Tyr Thr Arg Leu Arg Lys Gln  
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Met Ala Ala Lys Lys Tyr Leu Gln Ser Ile Lys Asn Lys Pro  
20 25 30

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<400> 101

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Met Ala Ala Lys Lys Tyr Leu Gln Ser Ile Lys Asn Lys Gln  
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His Ser Asp Ala Val Phe Thr Asp Gln Tyr Thr Arg Leu Arg Lys Gln  
1 5 10 15

Met Ala Ala Lys Lys Tyr Leu Gln Ser Ile Lys Asn Lys Ser  
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<400> 103

His Ser Asp Ala Val Phe Thr Asp Gln Tyr Thr Arg Leu Arg Lys Gln  
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Met Ala Ala Lys Lys Tyr Leu Gln Ser Ile Lys Asn Lys Thr  
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<400> 104

His Ser Asp Ala Val Phe Thr Asp Gln Tyr Thr Arg Leu Arg Lys Gln  
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Met Ala Ala Lys Lys Tyr Leu Gln Ser Ile Lys Asn Lys Val  
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<400> 105

His Ser Asp Ala Val Phe Thr Asp Gln Tyr Thr Arg Leu Arg Lys Gln  
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Met Ala Ala Lys Lys Tyr Leu Gln Ser Ile Lys Asn Lys Trp  
20 25 30

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His Ser Asp Ala Val Phe Thr Asp Gln Tyr Thr Arg Leu Arg Lys Gln  
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Met Ala Ala Lys Lys Tyr Leu Gln Ser Ile Lys Asn Lys Tyr  
20 25 30

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<400> 107

His Ser Asp Ala Val Phe Thr Asp Gln Tyr Thr Arg Leu Arg Lys Gln  
1 5 10 15

Met Ala Gly Lys Lys Tyr Leu Gln Ser Ile Lys Asn Arg Ile  
20 25 30

5189.ST25.txt

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<400> 108

His Ser Asp Ala Val Phe Thr Asp Gln Tyr Thr Arg Leu Arg Lys Gln  
1 5 10 15

Met Ala Lys Lys Tyr Leu Gln Ser Ile Lys Asn Arg Ile  
20 25 30

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His Ser Asp Ala Val Phe Thr Asp Gln Tyr Thr Arg Leu Arg Lys Gln  
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Met Ala Ser Lys Lys Tyr Leu Gln Ser Ile Lys Asn Arg Ile  
20 25 30

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<400> 110

His Ser Asp Ala Val Phe Thr Asp Gln Tyr Thr Arg Leu Arg Lys Gln  
1 5 10 15

Met Ala Ala Lys Lys Tyr Leu Gln Ser Ile Pro Asn Arg Ile  
20 25 30

5189.ST25.txt

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<400> 111

His Ser Asp Ala Val Phe Thr Asp Gln Tyr Thr Arg Leu Arg Lys Gln  
1 5 10 15

Met Ala Ser Lys Lys Tyr Leu Gln Ser Ile Arg Asn Arg Ile  
20 25 30

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<220>  
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<400> 112

His Ser Asp Ala Val Phe Thr Asp Gln Tyr Thr Arg Leu Arg Lys Gln  
1 5 10 15

Val Ala Ala Lys Lys Tyr Leu Gln Ser Ile Lys Gln Lys Arg Tyr Cys  
20 25 30

<210> 113  
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5189.ST25.txt

<223> ACETYLATION

<400> 113

His Ser Asp Ala Val Phe Thr Asp Gln Tyr Thr Arg Leu Arg Lys Gln  
1 5 10 15

Met Ala Ala Lys Lys Tyr Leu Gln Ser Ile Lys Gln Lys Arg Tyr Cys  
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<210> 114

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<223> ACETYLATION

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His Ser Asp Ala Val Phe Thr Asp Gln Tyr Thr Arg Leu Arg Lys Gln  
1 5 10 15

Val Ala Ala Lys Lys Tyr Leu Gln Ser Ile Lys Gln Lys Cys  
20 25 30

<210> 115

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<222> (1)..(32)

<223> ACETYLATION

<400> 115

His Thr Glu Ala Val Phe Thr Asp Gln Tyr Thr Arg Leu Arg Lys Gln  
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Val Ala Ala Lys Lys Tyr Leu Gln Ser Ile Lys Gln Lys Arg Tyr Cys  
20 25 30

5189.ST25.txt

<210> 116  
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<220>  
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<400> 116

His Ser Asp Ala Val Phe Thr Asp Gln Tyr Thr Arg Leu Arg Lys Gln  
1 5 10 15

Leu Ala Val Lys Lys Tyr Leu Gln Asp Ile Lys Gln Gly Gly Thr Cys  
20 25 30

<210> 117  
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<400> 117

His Ser Asp Ala Val Phe Thr Asp Gln Tyr Thr Arg Leu Arg Lys Gln  
1 5 10 15

Met Ala Ala Lys Lys Tyr Leu Gln Ser Ile Lys Gln Lys Arg Cys  
20 25 30

<210> 118  
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5189.ST25.txt

<220>  
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<400> 118

His Ser Asp Ala Val Phe Thr Asp Gln Tyr Thr Arg Leu Arg Lys Gln  
1 5 10 15

Leu Ala Ala Lys Lys Tyr Leu Gln Thr Ile Lys Gln Lys Arg Tyr Cys  
20 25 30

<210> 119  
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<400> 119

His Ser Asp Ala Val Phe Thr Asp Gln Tyr Thr Arg Leu Arg Lys Gln  
1 5 10 15

Met Ala Ala Lys Lys Tyr Leu Gln Thr Ile Lys Gln Lys Arg Tyr Cys  
20 25 30

<210> 120  
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His Ser Asp Ala Val Phe Thr Asp Gln Tyr Thr Arg Leu Arg Lys Gln  
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5189.ST25.txt

Met Ala Ala His Lys Tyr Leu Gln Ser Ile Lys Gln Lys Arg Tyr Cys  
20 25 30

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Met Ala Ala Lys His Tyr Leu Gln Ser Ile Lys Gln Lys Arg Tyr Cys  
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Met Ala Gly Lys Lys Tyr Leu Gln Ser Ile Lys Gln Lys Arg Cys  
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Met Ala Lys Lys Lys Tyr Leu Gln Ser Ile Lys Gln Lys Arg Cys  
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Met Ala Arg Lys Lys Tyr Leu Gln Ser Ile Lys Gln Lys Arg Cys  
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5189.ST25.txt

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Met Ala Ser Lys Lys Tyr Leu Gln Ser Ile Lys Gln Lys Arg Cys  
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1 5 10 15

Met Ala Ala Lys Lys Tyr Leu Gln Ser Ile Pro Gln Lys Arg Cys  
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Met Ala Ala Lys Lys Tyr Leu Gln Ser Ile Arg Gln Lys Arg Cys  
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Met Ala Ala Lys Lys Tyr Leu Gln Ser Ile Lys Gln Arg Arg Cys  
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Met Ala Ala Lys Lys Tyr Leu Gln Ser Ile Lys Gln Lys Ala Cys  
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Met Ala Ala Lys Lys Tyr Leu Gln Ser Ile Lys Gln Lys Phe Cys  
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Met Ala Ala Lys Lys Tyr Leu Gln Ser Ile Lys Gln Lys His Cys  
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5189.ST25.txt

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Met Ala Ala Lys Lys Tyr Leu Gln Ser Ile Lys Gln Lys Ile Cys  
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Met Ala Ala Lys Lys Tyr Leu Gln Ser Ile Lys Gln Lys Lys Cys  
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Met Ala Ala Lys Lys Tyr Leu Gln Ser Ile Lys Gln Lys Leu Cys  
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Met Ala Ala Lys Lys Tyr Leu Gln Ser Ile Lys Gln Lys Met Cys  
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5189.ST25.txt

Met Ala Ala Lys Lys Tyr Leu Gln Ser Ile Lys Gln Lys Pro Cys  
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Met Ala Ala Lys Lys Tyr Leu Gln Ser Ile Lys Gln Lys Gln Cys  
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Met Ala Ala Lys Lys Tyr Leu Gln Ser Ile Lys Gln Lys Ser Cys  
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Met Ala Ala Lys Lys Tyr Leu Gln Ser Ile Lys Gln Lys Val Cys  
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Met Ala Ala Lys Lys Tyr Leu Gln Ser Ile Lys Gln Lys Trp Cys  
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<400> 145

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Met Ala Lys Lys Lys Tyr Leu Gln Ser Ile Lys Gln Arg Ile Cys  
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Met Ala Ser Lys Lys Tyr Leu Gln Ser Ile Lys Gln Arg Ile Cys  
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Met Ala Ala Lys Lys Tyr Leu Gln Ser Ile Pro Gln Arg Ile Cys  
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<400> 148

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1 5 10 15

Met Ala Ser Lys Lys Tyr Leu Gln Ser Ile Arg Gln Arg Ile Cys  
20 25 30

<210> 149  
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<213> Homo sapiens

<400> 149

His Ser Asp Gly Ile Phe Thr Asp Ser Tyr Ser Arg Tyr Arg Lys Gln  
1 5 10 15

Met Ala Val Lys Lys Tyr Leu Ala Ala Val Leu Gly Lys Arg Tyr Lys  
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Gln Arg Val Lys Asn Lys  
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5189.ST25.txt

<400> 150

His Ser Asp Gly Ile Phe Thr Asp Ser Tyr Ser Arg Tyr Arg Lys Gln  
1 5 10 15

Met Ala Val Lys Lys Tyr Leu Ala Ala Val Leu  
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<210> 151

<211> 28

<212> PRT

<213> Homo sapiens

<400> 151

His Ser Asp Ala Val Phe Thr Asp Asn Tyr Thr Arg Leu Arg Lys Gln  
1 5 10 15

Met Ala Val Lys Lys Tyr Leu Asn Ser Ile Leu Asn  
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<210> 152

<211> 31

<212> PRT

<213> Homo sapiens

<400> 152

His Ser Asp Ala Val Phe Thr Asp Gln Tyr Thr Arg Leu Arg Lys Gln  
1 5 10 15

Val Ala Ala Lys Lys Tyr Leu Gln Ser Ile Lys Gln Lys Arg Tyr  
20 25 30

<210> 153

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<213> Homo sapiens

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<223> ACETYLATION

<400> 153

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Val Ala Ala Lys Lys Tyr Leu Gln Ser Ile Lys Gln Lys Arg Tyr  
20 25 30

<210> 154

5189.ST25.txt

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<400> 154

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1 5 10 15

Val Ala Ala Lys Lys Tyr Leu Gln Ser Ile Lys Gln Lys Arg Tyr Cys  
20 25 30

<210> 155  
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<400> 155

His Thr Asp Ala Val Phe Thr Asp Gln Tyr Thr Arg Leu Arg Lys Gln  
1 5 10 15

Val Ala Ala Lys Lys Tyr Leu Gln Ser Ile Lys Gln Lys Arg Tyr Cys  
20 25 30